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English The physiological role of circulating insulin-like growth factor-II

```
although HGP was similarly inhibited by insulin, phosphoenolpyruvate
     gluconeogenesis was enhanced and accounted for a larger portion of HGP
     (64% versus apprx 40% in control mice). Our data suggest that the
     persistence of circulating IGF-II in adult mice to levels commonly
     observed in adult humans (50-70 nm) causes a marked improvement in
     peripheral (skeletal muscle) insulin action, which is not due to
     changes in body composition. These results suggest that circulating IGF-II
     may exert a regulatory role on insulin sensitivity and body composition in
     humans.
     Genetics and Cytogenetics - Animal
     Clinical Biochemistry; General Methods and Applications
                                                                10006
     Biochemical Studies - Proteins, Peptides and Amino Acids
                                                                10064
     Biochemical Studies - Lipids
                                    10066
     Biochemical Studies - Carbohydrates
                                           10068
     Metabolism - General Metabolism; Metabolic Pathways *13002
     Metabolism - Carbohydrates *13004
     Metabolism - Lipids *13006
     Digestive System - Physiology and Biochemistry *14004
     Blood, Blood-Forming Organs and Body Fluids - Blood and Lymph Studies
     *15002
     Urinary System and External Secretions - Physiology and Biochemistry
     *15504
     Endocrine System - General *17002
     Endocrine System - Pancreas *17008
     Muscle - Physiology and Biochemistry *17504
    Muridae *86375
    Major Concepts
        Blood and Lymphatics (Transport and Circulation); Digestive System
        (Ingestion and Assimilation); Endocrine System (Chemical Coordination
        and Homeostasis); Metabolism; Muscular System (Movement and Support);
        Urinary System (Chemical Coordination and Homeostasis)
     Chemicals & Biochemicals
        INSULIN; FACTOR-II; GLUCOSE; LACTATE; PHOSPHOENOLPYRUVATE
     Miscellaneous Descriptors
        FREE FATTY ACID; GLUCOSE; GLUCOSE CLEARANCE; HEPATIC GLUCOSE
        PRODUCTION; INSULIN SENSITIVITY; LACTATE; PHOSPHOENOLPYRUVATE
        GLUCONEOGENESIS; PLASMA; SKELETAL MUSCLE INSULIN ACTION;
        TRANSGENIC MOUSE
ORGN Super Taxa
        Muridae: Rodentia, Mammalia, Vertebrata, Chordata, Animalia
ORGN Organism Name
        Muridae (Muridae)
ORGN Organism Superterms
        animals; chordates; mammals; nonhuman vertebrates; nonhuman mammals;
        rodents; vertebrates
     9004-10-8 (INSULIN)
     68-19-9Q (FACTOR-II)
     9001-26-7Q (FACTOR-II)
     50-99-7 (GLUCOSE)
     113-21-3 (LACTATE)
     73-89-2 (PHOSPHOENOLPYRUVATE)
    ANSWER 2 OF 4 CAPLUS COPYRIGHT 2001 ACS
     1996:41569 CAPLUS
     124:77363
    Hepatic overexpression of insulin-like growth factor-II in adulthood
     increases basal and insulin-stimulated glucose disposal in conscious mice
    Rossetti, Luciano; Barzilai, Nir; Chen, Wei; Harris, Thomas; Yang, Deyun;
     Rogler, Charles E.
     Division Endocrinology, Albert Einstein College Medicine, Bronx, NY,
     10461, USA
     J. Biol. Chem. (1996), 271(1), 203-8
    CODEN: JBCHA3; ISSN: 0021-9258
     Journal
     English
     2-10 (Mammalian Hormones)
    The physiol. role of circulating insulin-like growth factor-II (IGF-II) in
     adult humans is poorly understood. The authors recently generated an
     IGF-II transgenic murine model of persistent IGF-II prodn. (plasma IGF-II
     .apprx.30-fold increased above normal) through overexpression of the
     transgene driven by the major urinary protein
```

promoter. To det. whether in vivo insulin action is improved in these

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ΙΤ

ΙT

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exert a regulatory role on insulin sensitivity and body compn. in humans.
ST
     liver IGF II insulin glucose
IT
     Blood sugar
     Gluconeogenesis
     Glycolysis
     Liver
        (hepatic overexpression of IGF-II in adulthood increases basal and
        insulin-stimulated glucose disposal in conscious mice)
IT
     Biological transport
        (absorption, hepatic overexpression of IGF-II in adulthood increases
        basal and insulin-stimulated glucose disposal in conscious mice)
IT
     9004-10-8, Insulin, biological studies
     RL: BAC (Biological activity or effector, except adverse); BIOL
     (Biological study)
        (hepatic overexpression of IGF-II in adulthood increases basal and
        insulin-stimulated glucose disposal in conscious mice)
ΙT
     67763-97-7, IGF-II
     RL: BAC (Biological activity or effector, except adverse); MFM (Metabolic
     formation); BIOL (Biological study); FORM (Formation, nonpreparative)
        (hepatic overexpression of IGF-II in adulthood increases basal and
        insulin-stimulated glucose disposal in conscious mice)
     50-21-5, Lactic acid, biological studies
                                                50-99-7, D-Glucose, biological
ΙT
              138-08-9, Phosphoenolpyruvic acid
     studies
     RL: BPR (Biological process); BIOL (Biological study); PROC (Process)
        (hepatic overexpression of IGF-II in adulthood increases basal and
        insulin-stimulated glucose disposal in conscious mice)
ΙT
     9005-79-2, Glycogen, biological studies
     RL: MFM (Metabolic formation); BIOL (Biological study); FORM (Formation,
     nonpreparative)
        (hepatic overexpression of IGF-II in adulthood increases basal and
        insulin-stimulated glucose disposal in conscious mice)
    ANSWER 3 OF 4 EMBASE COPYRIGHT 2001 ELSEVIER SCI. B.V.
L5
     96023565 EMBASE
AN
    1996023565
DN
    Hepatic overexpression of insulin-like growth factor-II in adulthood
TI
     increases basal and insulin-stimulated glucose disposal in conscious mice.
     Rossetti L.; Barzilai N.; Chen W.; Harris T.; Yang D.; Rogler C.E.
ΑU
    Div. of Endocrinology, Dept. of Medicine, Albert Einstein College of
CS
    Medicine, 1300 Morris Park Ave., Bronx, NY 10461, United States
     Journal of Biological Chemistry, (1996) 271/1 (203-208).
SO
     ISSN: 0021-9258 CODEN: JBCHA3
CY
     United States
DT
     Journal; Article
FS
     003
            Endocrinology
     029
             Clinical Biochemistry
    English
LA
SL
     English
    The physiological role of circulating insulin-like growth factor-II (IGF-
AΒ
     II) in adult humans is poorly understood. We recently generated an IGF-II
     transgenic murine model of persistent IGF-II production (plasma IGF-II
     .apprx.30- fold increased above normal) through overexpression of the
     transgene driven by the major urinary protein
    promoter (Rinderknecht, E, and Humbel, R. E. (1978) J. Biol. Chem. 269,
     13779-13784). To determine whether in vivo insulin action is improved in
    these transgenic mice, we performed euglycemic insulin (18 milliunits/kg
     .cntdot. min) clamp studies in conscious IGF-II transgenic and in age- and
    weight-matched control mice. Plasma glucose and insulin concentrations
    were significantly lower in the IGF-II transgenic compared with both
     control groups. Despite decreased plasma glucose concentration, basal
    hepatic glucose production (HGP) and glucose clearance were increased.
     During the insulin clamp studies in IGF-II transgenic mice compared with
     control mice (a) the rates of glucose infusion and glucose uptake were
     increased by .apprx.65 and .apprx.55%, respectively; (b) glycolysis was
     increased by .apprx.12% while glycogen synthesis was .apprx.2-fold higher;
     (c) while the suppression of plasma free fatty acid was similar, the
     increment in plasma lactate concentration was significantly higher; (d)
     although HGP was similarly inhibited by insulin, phosphoenolpyruvate
     gluconeogenesis was enhanced and accounted for a larger portion of HGP
     (64% versus .apprx.40% in control mice). Our data suggest that the
```

persistence of circulating IGF-II in adult mice to levels commonly observed in adult humans $(50-70\ \mathrm{nM})$ causes a marked improvement in

```
insulin blood level
     mouse
     nonhuman
     priority journal
     promoter region
     transgene
     transgenic mouse
     Drug Descriptors:
     *insulin
     *somatomedin b: EC, endogenous compound
     glucose: EC, endogenous compound
     lactic acid: EC, endogenous compound
     (insulin) 9004-10-8; (somatomedin b) 63774-77-6, 67763-97-7; (glucose)
RN
     50-99-7, 84778-64-3; (lactic acid) 113-21-3, 50-21-5
L5
     ANSWER 4 OF 4 MEDLINE
AN
     96132904
                  MEDLINE
DN
     96132904
ΤI
     Hepatic overexpression of insulin-like growth factor-II in adulthood
     increases basal and insulin-stimulated glucose disposal in conscious mice.
ΑU
     Rossetti L; Barzilai N; Chen W; Harris T; Yang D; Rogler C E
CS
     Division of Endocrinology, Albert Einstein College of Medicine, Bronx, New
     York 10461, USA.
NC
     R029-DK 45024 (NIDDK)
     R01-DK 48321 (NIDDK)
     R01-CA 56076 (NCI)
     JOURNAL OF BIOLOGICAL CHEMISTRY, (1996 Jan 5) 271 (1) 203-8.
SO
     Journal code: HIV. ISSN: 0021-9258.
CY
     United States
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
FS
     Priority Journals; Cancer Journals
EM
     199604
     The physiological role of circulating insulin-like growth factor-II
AΒ
     (IGF-II) in adult humans is poorly understood. We recently generated an
     IGF-II transgenic murine model of persistent IGF-II production (plasma
     IGF-II approximately 30-fold increased above normal) through
     over-expression of the transgene driven by the major
     urinary protein promoter (Rinderknecht, E., and Humbel,
     R. E. (1978) J. Biol. Chem. 269, 13779-13784). To determine whether in
     vivo insulin action is improved in these transgenic mice, we performed
     euglycemic insulin (18 milliunits/kg.min) clamp studies in conscious
     IGF-II transgenic and in age- and weight-matched control mice. Plasma
     glucose and insulin concentrations were significantly lower in the IGF-II
     transgenic compared with both control grouoff Despite decreased plasma
     glucose concentration, basal hepatic glucose production (HGP) and glucose
     clearance were increased. During the insulin clamp studies in IGF-II
     transgenic mice compared with control mice (a) the rates of glucose
     infusion and glucose uptake were increased by approximately by 65 and
     approximately 55%, respectively; (b) glycolysis was increased by
     approximately 12% while glycogen synthesis was approximately 2-fold
     higher; (c) while the suppression of plasma free fatty acid was similar,
     the increment in plasma lactate concentration was significantly higher;
     (d) although HGP was similarly inhibited by insulin, phosphoenolpyruvate
     gluconeogenesis was enhanced and accounted for a larger portion of HGP
     (64% versus approximately 40% in control mice). Our data suggest that the
    persistence of circulating IGF-II in adult mice to levels commonly
     observed in adult humans (50-70 nM) causes a marked improvement in
     peripheral (skeletal muscle) insulin action, which is not due to
     changes in body composition. These results suggest that circulating IGF-II
    may exert a regulatory role on insulin sensitivity and body composition in
    Check Tags: Animal; Human; Support, Non-U.S. Gov't; Support, U.S. Gov't,
     Adult
     Glucokinase: ME, metabolism
     *Glucose: ME, metabolism
     Glucose Clamp Technique
      Glucose-6-Phosphatase: ME, metabolism
      Glycogen: ME, metabolism
     Glycogen Synthase: ME, metabolism
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CT

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     29 MAR 2001
L1
            169 S (ALPHA2U GLOBULIN)
              O S (MUSCLE TYPE FATTY ACID)
L2
L3
              0 S L1 AND MUSCLE?
L4
            743 S (MAJOR URINARY PROTEIN)
L5
              4 S L4 AND MUSCLE
=> s l1 and heart?
             4 L1 AND HEART?
=> d 16 1-4 all
L6
     ANSWER 1 OF 4 BIOSIS COPYRIGHT 2001 BIOSIS
     1999:505492 BIOSIS
ΑN
DN
     PREV199900505492
    Effects of a thirteen-week inhalation exposure to ethyl tertiary butyl
TΙ
     ether on Fischer-344 rats and CD-1 mice.
    Medinsky, M. A.; Wolf, D. C.; Cattley, R. C.; Wong, B. (1); Janszen, D.
ΑU
    B.; Farris, G. M.; Wright, G. A.; Bond, J. A.
     (1) Chemical Industry Institute of Toxicology, 6 Davis Drive, Research
CS
     Triangle Park, NC, 27709-2137 USA
     Toxicological Sciences, (Sept., 1999) Vol. 51, No. 1, pp. 108-118.
SO
     ISSN: 1096-6080.
DT
    Article
LA
    English
SL
    English
    The 1990 Clean Air Act Amendments require that oxygenates be added to
AΒ
     automotive fuels to reduce emissions of carbon monoxide and hydrocarbons.
    One potential oxygenate is the aliphatic ether ethyl tertiary butyl ether
     (ETBE). Our objective was to provide data on the potential toxic effects
     of ETBE. Male and female Fisher 344 rats and CD-1 mice were exposed to 0
     (control), 500, 1750, or 5000 ppm of ETBE for 6 h/day and 5 days/wk over a
     13-week period. ETBE exposure had no effect on mortality and body weight
    with the exception of an increase in body weights of the female rats in
    the 5000-ppm group. No major changes in clinical pathology parameters were
    noted for either rats or mice exposed to ETBE for 6 (rats only) or 13
    weeks. Liver weights increased with increasing ETBE-exposure concentration
    for both sexes of rats and mice. Increases in kidney, adrenal, and
    heart (females only) weights were noted in rats. Degenerative
    changes in testicular seminiferous tubules were observed in male rats
    exposed to 1750 and 5000 ppm but were not seen in mice. This testicular
    lesion has not been reported previously for aliphatic ethers. Increases in
    the incidence of regenerative foci, rates of renal cell proliferation, and
    alpha2u-globulin containing protein droplets were noted
    in the kidneys of all treated male rats. These lesions are associated with
    the male rat-specific syndrome of alpha2u-globulin
    nephropathy. Increases in the incidence of centrilobular hepatocyte
    hypertrophy and rates of hepatocyte cell proliferation were seen in the
    livers of male and female mice in the 5000-ppm group, consistent with a
    mitogenic response to ETBE. These two target organs for ETBE toxicity,
    mouse liver and male rat kidney, have also been reported for methyl
    tertiary butyl ether and unleaded gasoline.
    Toxicology - General; Methods and Experimental *22501
CC
    Cytology and Cytochemistry - Animal *02506
    Digestive System - General; Methods *14001
    Cardiovascular System - General; Methods *14501
    Toxicology - Environmental and Industrial Toxicology *22506
    Public Health: Environmental Health - Air, Water and Soil Pollution
     *37015
    Urinary System and External Secretions - General; Methods *15501
    Reproductive System - General; Methods *16501
    Endocrine System - General *17002
    Muridae 86375
BC
```

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Animals; Chordates; Mammals; Nonhuman Mammals; Nonhuman Vertebrates;
        Rodents; Vertebrates
L6
     ANSWER 2 OF 4 BIOSIS COPYRIGHT 2001 BIOSIS
AN
     1998:221180 BIOSIS
DN
     PREV199800221180
ТΙ
     alpha2u-Globulin is not a bona fide fatty acid-binding
     protein in the rat kidney.
ΑU
     Khan, K. M. Faisal; Sato, Atsushi; Oka, Tatsuzo; Horiuchi, Saburou;
     Tsugita, Akira; Natori, Yasuo (1)
CS
     (1) Dep. Nutritional Chem., Sch. Med., Univ. Tokushima, Kuramoto,
     Tokushima 770 Japan
SO
     Research Communications in Biochemistry and Cell & Molecular Biology,
     (1997) Vol. 1, No. 1, pp. 33-42.
     ISSN: 1087-111X.
DT
     Article
LA
     English
AΒ
     It has been reported that the cytosolic fraction of male rat kidneys
     contains two different fatty acid-binding proteins (FABPs); one is
     identical to heart FABP and the other is proteolytically
     modified form of alpha2u-globulin. Highly purified
     lysosomes were isolated from the male rat kidney and subjected to
     immunoblot analysis using antibody against alpha2u-
     globulin. A major immunoreactive band was observed at the position
     about 1-kDa smaller than that of the authentic {\tt alpha2u}-
     globulin. Amino acid sequence analysis of the immuno-reactive
     protein established that the protein represented proteolytically modified
     alpha2u-globulin, lacking N-terminal 9 amino acid
     residues. The modified alpha2u-globulin was found to
     be the most abundant protein in the lysosomes, amounting to as much as 21%
     of the total lysosomal proteins. The occurrence of alpha2u-
     globulin in the cytosolic fraction, reported by earlier workers,
     was shown to be an artifact of isolation procedure. Since FABP must be
     cytosolic in order to be involved in intracellular transport and
     metabolism of fatty acids, alpha2u-globulin can not be
     considered as a bonafide member of the FABP superfamily.
CC
     Urinary System and External Secretions - Physiology and Biochemistry
     *15504
     Cytology and Cytochemistry - Animal *02506
     Biochemical Studies - Proteins, Peptides and Amino Acids *10064
     Biochemical Studies - Lipids *10066
BC
     Muridae
               86375
ΙT
     Major Concepts
        Biochemistry and Molecular Biophysics; Urinary System (Chemical
        Coordination and Homeostasis)
     Parts, Structures, & Systems of Organisms
ΙT
        kidney: excretory system
ΙT
     Chemicals & Biochemicals
        alpha-2-u-globulin; fatty acid-binding proteins: cytosolic, lysosomal
ORGN Super Taxa
        Muridae: Rodentia, Mammalia, Vertebrata, Chordata, Animalia
ORGN Organism Name
        rat (Muridae)
ORGN Organism Superterms
        Animals; Chordates; Mammals; Nonhuman Mammals; Nonhuman Vertebrates;
        Rodents; Vertebrates
     ANSWER 3 OF 4 CAPLUS COPYRIGHT 2001 ACS
L6
AN
     1997:45898 CAPLUS
DN
     126:128429
ΤI
     .alpha.2u-Globulin is not a bona fide fatty acid-binding protein in the
     rat kidney
ΑU
     Faisal Khan, K. M.; Sato, Atsushi; Oka, Tatsuzo; Horiuchi, Saburou;
     Tsugita, Akira; Natori, Yasuo
CS
     Sch. Med., Univ. Tokushima, Tokushima, 770, Japan
     Res. Commun. Biochem. Cell Mol. Biol. (1997), 1(1), 33-42
SO
     CODEN: RCBBFC; ISSN: 1087-111X
PB
     PJD Publications
DT
     Journal
LA
     English
CC
     6-3 (General Biochemistry)
     Section cross-reference(s): 13
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     The hear house on
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Lysosome
        (alpha2u-globulin is not a bona fide fatty
        acid-binding protein in the rat kidney)
ΙT
     Fatty acid-binding protein
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PRP
     (Properties); BIOL (Biological study); OCCU (Occurrence)
        (alpha2u-globulin is not a bona fide fatty
        acid-binding protein in the rat kidney)
ΙT
     Globulins, biological studies
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PRP
     (Properties); BIOL (Biological study); OCCU (Occurrence)
        (.alpha.2u-globulin; alpha2u-globulin is not a bona
        fide fatty acid-binding protein in the rat kidney)
L6
     ANSWER 4 OF 4 MEDLINE
ΑN
     1999425001
                    MEDLINE
DN
     99425001
ΤI
     Effects of a thirteen-week inhalation exposure to ethyl tertiary butyl
     ether on fischer-344 rats and CD-1 mice.
     Medinsky M A; Wolf D C; Cattley R C; Wong B; Janszen D B; Farris G M;
AU
     Wright G A; Bond J A
     Chemical Industry Institute of Toxicology, Research Triangle Park, North
CS
     Carolina 27709-2137, USA.
     TOXICOLOGICAL SCIENCES, (1999 Sep) 51 (1) 108-18.
SO
     Journal code: CZ1. ISSN: 1096-6080.
CY
     United States
     Journal; Article; (JOURNAL ARTICLE)
DT
LA
     English
FS
     Priority Journals
EM
     199912
ΕW
     19991204
AΒ
     The 1990 Clean Air Act Amendments require that oxygenates be added to
     automotive fuels to reduce emissions of carbon monoxide and hydrocarbons.
     One potential oxygenate is the aliphatic ether ethyl tertiary butyl ether
     (ETBE). Our objective was to provide data on the potential toxic effects
     of ETBE. Male and female Fisher 344 rats and CD-1 mice were exposed to 0
     (control), 500, 1750, or 5000 ppm of ETBE for 6 h/day and 5 days/wk over a
     13-week period. ETBE exposure had no effect on mortality and body weight
     with the exception of an increase in body weights of the female rats in
     the 5000-ppm group. No major changes in clinical pathology parameters were
     noted for either rats or mice exposed to ETBE for 6 (rats only) or 13
     weeks. Liver weights increased with increasing ETBE-exposure concentration
     for both sexes of rats and mice. Increases in kidney, adrenal, and
     heart (females only) weights were noted in rats. Degenerative
     changes in testicular seminiferous tubules were observed in male rats
     exposed to 1750 and 5000 ppm but were not seen in mice. This testicular
     lesion has not been reported previously for aliphatic ethers. Increases in
     the incidence of regenerative foci, rates of renal cell proliferation, and
     alpha2u-globulin containing protein droplets were noted
     in the kidneys of all treated male rats. These lesions are associated with
     the male rat-specific syndrome of alpha2u-globulin
     nephropathy. Increases in the incidence of centrilobular hepatocyte
     hypertrophy and rates of hepatocyte cell proliferation were seen in the
     livers of male and female mice in the 5000-ppm group, consistent with a
     mitogenic response to ETBE. These two target organs for ETBE toxicity,
     mouse liver and male rat kidney, have also been reported for methyl
     tertiary butyl ether and unleaded gasoline.
     Check Tags: Animal; Comparative Study; Female; Male; Support, Non-U.S.
CT
     Gov't
      Administration, Inhalation
     *Air Pollutants, Environmental: TO, toxicity
      Alpha-Globulins: ME, metabolism
      Atmosphere Exposure Chambers
      Body Weight: DE, drug effects
      Bone Marrow: DE, drug effects
      Bone Marrow: PA, pathology
      Bromodeoxyuridine: ME, metabolism
      Cell Division: DE, drug effects
     *Ethyl Ethers: TO, toxicity
      Kidney: DE, drug effects
      Kidney: ME, metabolism
      Kidney: PA, pathology
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ΑN

DN

2000:497084 CAPLUS

133:191444

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     29 MAR 2001
Ll
            169 S (ALPHA2U GLOBULIN)
L2
              O S (MUSCLE TYPE FATTY ACID)
L3
              0 S L1 AND MUSCLE?
L4
            743 S (MAJOR URINARY PROTEIN)
L5
              4 S L4 AND MUSCLE
L6
              4 S L1 AND HEART?
=> s 14 and heart
             7 L4 AND HEART
T.7
=> d 17 1-7 all
     ANSWER 1 OF 7 BIOSIS COPYRIGHT 2001 BIOSIS
L7
AN
     1991:234244
                 BIOSIS
DN
     BA91:125704
     PRIMARY STRUCTURE AND CELLULAR DISTRIBUTION OF TWO FATTY ACID-BINDING
TI
     PROTEINS IN ADULT RAT KIDNEYS.
ΑU
     KIMURA H; ODANI S; NISHI S-I; SATO H; ARAKAWA M; ONO T
     DEP. BIOCHEM., NIIGATA UNIV. SCH. MED., NIIGATA 951, JPN.
CS
     J BIOL CHEM, (1991) 266 (9), 5963-5972.
SO
     CODEN: JBCHA3. ISSN: 0021-9258.
FS
     BA; OLD
LA
     English
AB
     Fatty acid-binding proteins (FABPs) were purified from the kidneys of
     female and male rats and characterized by primary structure and
     histological distribution in the kidney. Two FABPs (14 and 15.5 kDa) were
     found in male rat kidney cytosol whereas only 14-kDa FABP could be
     recognized in female rat kidneys throughout the purification steps. The
     amino acid sequence of the 14-kDa FABP was identical to that of rat
    heart FABP deduced from the cDNA sequence (Heuckeroth, R. O.,
    Birkenmeier, E. H., Levin, M. S., and Gordon, J. I. (1987) J. Biol. Chem.
     262, 9709-9717). Structural analysis of the male-specific 15.5-kDa FABP
     identified this second FABP as a proteolytically modified form of
     .alpha.2u-globulin, an 18.7-kDa major urinary
    protein of adult male rats (Unterman, R. D., Lynch, K. R.,
    Nakhasi, H. L., Dolan, K. P., Hamilton, J. W., Cohn, D. V., and Feigelson,
     P. (1981) Proc. Natl. Acad. Sci. U.S.A. 78, 3478-3482) which shares a
     common ancestry with a number of hydrophobic ligand-binding proteins such
     as serum retinol-binding proteins. Immunohistochemical investigation
    disclosed that heart-type FABP (14-kDa FABP) is localized in the
    cytoplasm of the epithelia of the distal tubules in both male and female
    rat kidneys whereas 15.5-kDa FABP immunostaining was observed
    predominantly in the endosomes or lysosomes of proximal tubules in male
    rat kidneys. These results suggest strongly the functional divergence of
    two FABPs in the rat kidney.
    Microscopy Techniques - Cytology and Cytochemistry 01054
    Cytology and Cytochemistry - Animal *02506
    Genetics and Cytogenetics - Sex Differences
    Biochemical Studies - Proteins, Peptides and Amino Acids 10064
    Biophysics - Molecular Properties and Macromolecules 10506
    Anatomy and Histology, General and Comparative - Microscopic and
    Ultramicroscopic Anatomy *11108
    Urinary System and External Secretions - Physiology and Biochemistry
     *15504
BC
    Muridae 86375
IT
    Miscellaneous Descriptors
        SEX DIFFERENCES HISTOLOGY MOLECULAR SEQUENCE DATA AMINO ACID SEQUENCE
    ANSWER 2 OF 7 CAPLUS COPYRIGHT 2001 ACS
L7
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signal-transduction pathway. Here we use mice unable to express SOCS-2 to
    examine its function in vivo. SOCS-2-/- mice grew significantly larger
    than their wild-type littermates. Increased body wt. became evident after
    weaning and was assocd. with significantly increased long bone lengths and
     the proportionate enlargement of most organs. Characteristics of
    deregulated growth hormone and insulin-like growth factor-I (IGF-I)
    signaling, including decreased prodn. of major urinary
    protein, increased local IGF-I prodn., and collagen accumulation
    in the dermis, were obsd. in SOCS-2-deficient mice, indicating that SOCS-2
    may have an essential neg. regulatory role in the growth hormone/IGF-I
    pathway.
    SOCS2 protein deficiency gigantism
    Proteins, specific or class
    RL: BOC (Biological occurrence); BIOL (Biological study); OCCU
     (Occurrence)
        (MUP (major urinary protein); gigantism
        in mice lacking suppressor of cytokine signaling-2)
     Proteins, specific or class
    RL: BAC (Biological activity or effector, except adverse); BOC (Biological
     occurrence); BIOL (Biological study); OCCU (Occurrence)
        (SOCS-2 (suppressor of cytokine signaling-2); gigantism in mice lacking
        suppressor of cytokine signaling-2)
    Skin
        (dermis, collagen accumulation in; gigantism in mice lacking suppressor
        of cytokine signaling-2)
     Sex differences
    Signal transduction, biological
        (qiqantism in mice lacking suppressor of cytokine signaling-2)
    Growth disorders, animal
        (gigantism; gigantism in mice lacking suppressor of cytokine
        signaling-2)
    Adipose tissue
    Bladder
    Heart
    Liver
    Lung
     Testis
        (growth in wt. of; gigantism in mice lacking suppressor of cytokine
        signaling-2)
    Collagens, biological studies
    RL: BOC (Biological occurrence); BIOL (Biological study); OCCU
     (Occurrence)
        (in dermis; gigantism in mice lacking suppressor of cytokine
        signaling-2)
                                 67763-96-6, Insulin-like growth factor I
     9002-72-6, Growth hormone
     RL: BAC (Biological activity or effector, except adverse); BIOL
     (Biological study)
        (gigantism in mice lacking suppressor of cytokine signaling-2)
RE.CNT
(1) Adams, T; J Biol Chem 1998, V273, P1285 CAPLUS
(2) Alexander, W; Blood 1996, V87, P2162 CAPLUS
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(22) Starr, R; Proc Natl Acad Sci USA 1998, V95, P14395 CAPLUS
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מין יפשר במצבת האינו מססר נויה ביה ביה ביה

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FAN.CNT 1
     PATENT NO.
                     KIND DATE
                                           APPLICATION NO. DATE
                     A1 19990603
PΙ
     WO 9927363
                                           WO 1998-JP5319 19981126
        W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IS, KE, KG,
             KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO,
             NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA,
             UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
             FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
             CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                                            19981124
                     A2 19990907
                                           JP 1998-331828
     JP 11242026
                                           AU 1999-12603
     AU 9912603
                       Α1
                            19990615
                                                            19981126
                                           EP 1998-955936
                                                            19981126
     EP 1043587
                       Α1
                            20001011
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
PRAI JP 1997-323684
                      19971126
     WO 1998-JP5319
                      19981126
    A diagnostic method is described for examg. kidney diseases by immunol.
AB
     detecting a fatty acid-binding protein derived from kidney tissues
     contained in the specimen sampled from mammals other than rodents.
     method can provide examn. results contg. information highly useful in
     diagnosing the prognosis of kidney diseases hardly obtained by the
     existing methods. Based on the results obtained by this method, an
     appropriate therapy can be selected by taking the risk concerning the
     prognosis into consideration. This method is applicable not only to
     kidney tissue samples, but also to urine samples, and therefore, the
     examn. can be conveniently and efficiently performed.
     kidney disease diagnosis prognosis immunoassay staining; fatty acid
ST
     binding protein renal failure
ΙT
     Fatty acid-binding protein
     RL: ANT (Analyte); BUU (Biological use, unclassified); PUR (Purification
     or recovery); THU (Therapeutic use); ANST (Analytical study); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (L-FABP (liver fatty acid-binding protein), human, mouse, rabbit;
        method for examg. kidney diseases)
ΙT
     Proteins (specific proteins and subclasses)
     RL: ANT (Analyte); THU (Therapeutic use); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
        (MUP (major urinary protein); method for
        examq. kidney diseases)
ΙT
    Antibodies
     RL: ARG (Analytical reagent use); BPN (Biosynthetic preparation); ANST
     (Analytical study); BIOL (Biological study); PREP (Preparation); USES
        (anti-mouse L-FABP, anti-mouse H-FABP, anti-human L-FABP,; method for
        examg. kidney diseases)
    Basement membrane
TΤ
        (glomerular; method for examg. kidney diseases)
ΙT
     Phosphoproteins
     RL: ANT (Analyte); BUU (Biological use, unclassified); PUR (Purification
     or recovery); THU (Therapeutic use); ANST (Analytical study); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (h-FABP (heart fatty acid-binding protein), mouse; method for
        examg. kidney diseases)
ΙT
     Renal fibrosis
        (interstitial; method for examg. kidney diseases)
IT
     Blood analysis
     Diagnosis
     Disease models
     Distal tubule (kidney)
     ELISA (immunosorbent assay)
     IgA nephropathy
     Immunological staining
     Kidney
     Kidney diseases
    Mammal (Mammalia)
    Mouse
     Polyacrylamide gel electrophoresis
     Prognosis
     Proximal tubule (kidney)
```

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(2) Uchida; Febs Lett 1995, V357(2), P165 CAPLUS
     ANSWER 4 OF 7 CAPLUS COPYRIGHT 2001 ACS
L7
ΑN
     1997:357084 CAPLUS
DN
     127:63583
TΤ
     Epigenetic inheritance in the mouse
ΆU
     Roemer, Irmgard; Reik, Wolf; Dean, Wendy; Klose, Joachim
     Institut Humangenetik, Humboldt Universitat, Institut Toxikologie
CS
     Embryopharmakologie, Freie Universitat, Berlin, D-14195, Germany
     Curr. Biol. (1997), 7(4), 277-280
SO
     CODEN: CUBLE2; ISSN: 0960-9822
PΒ
     Current Biology
DT
     Journal
LA
     English
CC
     13-3 (Mammalian Biochemistry)
     Section cross-reference(s): 3
     Acquired epigenetic modifications, such as DNA methylation or stable
AB
     chromatin structures, are not normally thought to be inherited through the
     germline to future generations in mammals. Studies in the mouse have
     shown that specific manipulations of early embryos, such as nuclear
     transplantation, can result in altered patterns of gene expression and
     induce phenotypic alterations at later stages of development.
     effects are consistent with acquired epigenetic modifications that are
     somatically heritable, such as DNA methylation. Repression and DNA
     methylation of genes encoding major urinary
     proteins, repression of the gene encoding olfactory marker
     protein, and reduced body wt. can be exptl. induced by nuclear
     transplantation in early embryos. Strikingly, we now report that these acquired phenotypes are transmitted to most of the offspring of
     manipulated parent mice. This is the first demonstration of epigenetic
     inheritance of specific alterations of gene expression through the
     germline. These observations establish a mammalian model for
     transgenerational effects that are important for human health, and also
     raise the question of the evolutionary importance of epigenetic
     inheritance.
ST
     epigenetic inheritance DNA methylation gene expression
ΙT
     Genes (animal)
     RL: BPR (Biological process); BIOL (Biological study); PROC (Process)
        (Mup, for major urinary proteins;
        epigenetic inheritance in mouse)
IT
     Genes (animal)
     RL: BPR (Biological process); BIOL (Biological study); PROC (Process)
        (Omp, for olfactory marker protein; epigenetic inheritance in mouse)
ΙT
     Embryo (animal)
        (early 1-5 all; epigenetic inheritance in mouse)
ΙT
     Brain
     DNA methylation
     Gene expression
     Heart
     Inheritance (genetic)
     Liver
        (epigenetic inheritance in mouse)
ΙT
        (imprinting; epigenetic inheritance in mouse)
ΙT
     Transformation (genetic method)
        (mouse; epigenetic inheritance in mouse)
     Body weight
ΙT
        (reduced; epigenetic inheritance in mouse)
L7
     ANSWER 5 OF 7 CAPLUS COPYRIGHT 2001 ACS
ΑN
     1991:577629 CAPLUS
DN
     115:177629
     Primary structure and cellular distribution of two fatty acid-binding
TI
     proteins in adult rat kidneys
     Kimura, Hideki; Odani, Shoji; Nishi, Shinichi; Sato, Hirokazu; Arakawa,
ΑU
     Masaaki; Ono, Teruo
     Sch. Med., Niigata Univ., Niigata, 951, Japan
CS
     J. Biol. Chem. (1991), 266(9), 5963-72
SO
     CODEN: JBCHA3; ISSN: 0021-9258
DT
     Journal
LA
     English
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2 3 1000

יווי בשני שני שני שני שני בי יווי

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These results suggest strongly the functional divergence of two FABPs in
     the rat kidney.
ST
     fatty acid binding protein sequence kidney; kidney FABP cellular
     distribution
ΙT
     Kidney, composition
        (fatty acid-binding protein 14,000-mol.-wt. and 15,500-mol-wt. forms
        of, amino acid sequence and cellular distribution of)
ΙT
     Heart, composition
        (fatty acid-binding protein 14,000-mol.-wt. protein of, protein of
        kidney identity with)
ΙT
        (fatty acid-binding proteins cellular distribution in kidney in
        relation to)
ΙT
     Protein sequences
        (of fatty acid-binding protein 14,000-mol.-wt. form, of kidney,
ΙT
     Disulfide group
        (of fatty acid-binding protein 15,500-mol.-wt. form of kidney,
        localization of)
ΙT
     Protein sequences
        (of fatty acid-binding protein 15,500-mol.-wt. form, of kidney,
        complete)
ΙT
     Proteins, specific or class
     RL: BIOL (Biological study)
        (FABP (fatty acid-binding protein), kidney, 14,000-mol.-wt., amino acid
        sequence and cellular distribution of, sex in relation to)
ΙT
     Proteins, specific or class
     RL: BIOL (Biological study)
        (FABP (fatty acid-binding protein), kidney, 15,500-mol.-wt., amino acid
        sequence and cellular distribution of, sex in relation to)
ΙT
     Globulins, properties
     RL: PRP (Properties)
        (.alpha.2u-, fatty acid-binding protein 15,500-mol.-wt. form of kidney
        amino acid sequence homol. with)
     78849-32-8, .alpha.2u-Globulin (rat liver protein moiety reduced)
ΙT
     136602-37-4, .alpha.2u-Globulin (rat kidney)
                                                   136626-46-5, Protein FABP
     (rat kidney 14.0-kilodalton)
     RL: PRP (Properties)
        (amino acid sequence of)
L7
     ANSWER 6 OF 7 EMBASE COPYRIGHT 2001 ELSEVIER SCI. B.V.
ΑN
     91303044 EMBASE
     1991303044
DN
ΤI
     Primary structure and cellular distribution of two fatty acid-binding
     proteins in adult rat kidneys.
ΑU
     Kimura H.; Odani S.; Nishi S.-I.; Sato H.; Arakawa M.; Ono T.
CS
     Dept. of Biochemistry, Niigata University, School of Medicine, 1-757
     Asahimachi-dori, Niigata 951, Japan
     Journal of Biological Chemistry, (1991) 266/9 (5963-5972).
SO
     ISSN: 0021-9258 CODEN: JBCHA3
CY
     United States
DT
     Journal; Article
FS
     029
             Clinical Biochemistry
LA
     English
SL
     English
     Fatty acid-binding proteins (FABPs) were purified from the kidneys of
AΒ
     female and male rats and characterized by primary structure and
     histological distribution in the kidney. Two FABPs (14 and 15.5 kDa) were
     found in male rat kidney cytosol whereas only 14-kDa FABP could be
     recognized in female rat kidneys throughout the purification steps. The
     amino acid sequence of the 14-kDa FABP was identical to that of rat
     heart FABP deduced from the cDNA sequence (Heuckeroth, R.O.,
     Birkenmeier, E.H., Levin, M.S., and Gordon, J.I. (1987) J. Biol. Chem.
     262, 9709-9717). Structural analysis of the male-specific 15.5-kDa FABP
     identified this second FABP as a proteolytically modified form of
     .alpha.(2u)-globulin, an 18.7-kDa major urinary
     protein of adult male rats (Unterman, R.D., Lynch, K.R., Nakhasi,
     H.L., Dolan, K.P., Hamilton, J.W., Cohn, D.V., and Feigelson, P. (1981)
     Proc. Natl. Acad. Sci. U.S.A. 78, 3478-3482) which shares a common
     ancestry with a number of hydrophobic ligand-binding proteins such as
     serum retinol-binding proteins. Immunohistochemical investigation
     disclosed that heart-type FABP (14-kDa FABP) is localized in the
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of the emithelia of the distal tubules in both male and female

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L7
     ANSWER 7 OF 7 MEDLINE
AN
     91170283
                  MEDLINE
DN
     91170283
     Primary structure and cellular distribution of two fatty acid-binding
ΤI
     proteins in adult rat kidneys.
     Kimura H; Odani S; Nishi S; Sato H; Arakawa M; Ono T
ΑU
     Department of Biochemistry, Niigata University School of Medicine, Japan..
CS
SO
     JOURNAL OF BIOLOGICAL CHEMISTRY, (1991 Mar 25) 266 (9) 5963-72.
     Journal code: HIV. ISSN: 0021-9258.
CY
     United States
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
FS
     Priority Journals; Cancer Journals
ΕM
     199106
AΒ
     Fatty acid-binding proteins (FABPs) were purified from the kidneys of
     female and male rats and characterized by primary structure and
     histological distribution in the kidney. Two FABPs (14 and 15.5 kDa) were
     found in male rat kidney cytosol whereas only 14-kDa FABP could be
     recognized in female rat kidneys throughout the purification steps. The
     amino acid sequence of the 14-kDa FABP was identical to that of rat
     heart FABP deduced from the cDNA sequence (Heuckeroth, R. O.,
     Birkenmeier, E. H., Levin, M. S., and Gordon, J. I. (1987) J. Biol. Chem.
     262, 9709-9717). Structural analysis of the male-specific 15.5-kDa FABP
     identified this second FABP as a proteolytically modified form of alpha
     2u-globulin, an 18.7-kDa major urinary protein
     of adult male rats (Unterman, R. D., Lynch, K. R., Nakhasi, H. L., dolan,
     K. P., Hamilton, J. W., Cohn, D. V., and Feigelson, P. (1981) Proc. Natl.
     Acad. Sci. U.S.A. 78, 3478-3482) which shares a common ancestry with a
     number of hydrophobic ligand-binding proteins such as serum
     retinol-binding proteins. Immunohistochemical investigation disclosed that
     heart-type FABP (14-kDa FABP) is localized in the cytoplasm of the
     epithelia of the distal tubules in both male and female rat kidneys
     whereas 15.5-kDa FABP immunostaining was observed predominantly in the
     endosomes or lysosomes of proximal tubules in male rat kidneys. These
     results suggest strongly the functional divergence of two FABPs in the rat
CT
     Check Tags: Animal; Female; Male; Support, Non-U.S. Gov't
      Amino Acid Sequence
     *Carrier Proteins: ME, metabolism
      Chromatography, Gel
      Chromatography, High Pressure Liquid
      Electrophoresis, Polyacrylamide Gel
      Immunohistochemistry
     *Kidney: ME, metabolism
      Kidney: UL, ultrastructure
      Microscopy, Electron
      Molecular Sequence Data
      Myocardium: ME, metabolism
      Rats
```

Sequence Alignment

0 (fatty acid-binding proteins); 0 (Carrier Proteins)

Sex Factors

CN

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L4 ANSWER 7 OF 7 MEDLINE
AN 90194139 MEDLINE
DN 90194139
```

TI A comparison of male rat and **human** urinary proteins: implications for **human** resistance to hyaline droplet nephropathy.

AU Olson M J; Johnson J T; Reidy C A

CS Biomedical Science Department, General Motors Research Laboratories,
 Warren, Michigan 48090..
SO TOXICOLOGY AND APPLIED PHARMACOLOGY, (1990 Mar 1) 102 (3) 524-36.

· TOXICOLOGY AND APPLIED PHARMACOLOGY, (1990 Mar 1) 102 (3) 524-36. Journal code: VWO. ISSN: 0041-008X.

CY United States

DT Journal; Article; (JOURNAL ARTICLE)

LA English

FS Priority Journals; Cancer Journals

EM 199006

AΒ

alpha 2u-Globulin (alpha G), the major urinary protein of sexually mature male rats, is a key determinant of susceptibility to hyaline droplet nephropathy (HDN) induced by a variety of hydrocarbons in male rats. Arguments against extrapolating renal toxicity and carcinogenicity data for HDN-inducing toxicants from male rats to risk assessment for humans rely on the observation that humans do not express alpha G. Yet, human serum and urine are known to contain proteins coded for by the same gene family that also controls alpha G synthesis in the rat. Therefore, to understand some of the quantitative and qualitative differences between proteins of human and male rat urine which confer apparent resistance to HDN in humans, urinary proteins of male F344 rats (ca. 3 months old) and normal human males were compared by cation exchange, gel filtration, SDS-PAGE, and partially identified by Western blotting. We observed that (1) the protein content of human urine is only 1% that of male rat urine; (2) human urinary proteins, recovered by (NH4)2SO4 precipitation followed by dialysis, are primarily of high (greater than or equal to 75 kDa) molecular weight (MW) with minor components of 12-66 kDa; (3) male rat urine has little high-MW protein, but is rich in alpha G (18.5 kDa); (4) at pH 5, the most cationic fraction of human urinary protein constituted only about 4% of the total while the analogous fraction of rat urine, containing alpha G, contained 26% of total urinary protein; and (5) cationic (at pH 5.0) human urinary proteins included small amounts of proteins, e.g., alpha 1-acid glycoprotein, and alpha 1-microglobulin, which are products of the gene family coding for alpha G in rat. Thus, although humans excrete trace amounts of proteins similar to alpha G, the very low protein content of human urine, the relatively small proportion of cationic to total proteins, and the high ${\tt MW}$ of the most abundant ${\tt human}$ urinary proteins form a biological basis for suggesting that humans are not at risk for the type of fuel and solvent hydrocarbon-induced nephropathy, and the sequelae of such nephropathy,

CT Check Tags: Animal; Comparative Study; Human; Male Adult

Chromatography, Gel

observed in male rats.

Chromatography, Ion Exchange

Electrophoresis, Polyacrylamide Gel

*Hyalin: ME, metabolism

*Kidney Diseases: CI, chemically induced

Kidney Diseases: UR, urine
*Lysosomes: ME, metabolism

Molecular Weight

*Proteinuria: CI, chemically induced

Proteinuria: UR, urine

Rats

```
ANSWER 4 OF 7 EMBASE COPYRIGHT 2001 ELSEVIER SCI. B.V.
ΑN
     90097029 EMBASE
     1990097029
DN
TI
     A comparison of male rat and human urinary proteins:
     Implications for human resistance to hyaline droplet
     nephropathy.
     Olson M.J.; Johnson J.T.; Reidy C.A.
ΑIJ
CS
     Biomedical Science Department, General Motors Research Laboratories,
     Warren, MI 48090, United States
     Toxicology and Applied Pharmacology, (1990) 102/3 (524-536).
SO
     ISSN: 0041-008X CODEN: TXAPA
CY
     United States
DT
     Journal; Article
FS
     028
             Urology and Nephrology
     029
             Clinical Biochemistry
             Occupational Health and Industrial Medicine
     035
     052
             Toxicology
LA
    English
SL
     English
     .alpha.(2u)-Globulin (.alpha.G), the major
AB
     urinary protein of sexually mature male rats, is a key ·
     determinant of susceptibility to hyaline droplet nephropathy (HDN) induced
     by a variety of hydrocarbons in male rats. Arguments against extrapolating
     renal toxicity and carcinogenicity data for HDN-inducing toxicants from
     male rats to risk assessment for humans rely on the observation
     that humans do not express .alpha.G. Yet, human serum
     and urine are known to contain proteins coded for by the same gene family
     that also controls .alpha.G synthesis in the rat. Therefore, to understand
     some of the quantitative and qualitative differences between proteins of
     human and male rat urine which confer apparent resistance to HDN
     in humans, urinary proteins of male F344 rats (ca. 3 months old)
     and normal human males were compared by cation exchange, gel
     filtration, SDS-PAGE, and partially identified by Western blotting. We
     observed that (1) the protein content of human urine is only 1%
     that of male rat urine; (2) human urinary proteins, recovered by
     (NH4)2SO4 precipitation followed by dialysis, are primarily of
     high(.gtoreq.75 kDa) molecular weight (MW) with minor components of 12-66
     kDa; (3) male rat urine has little high-MW protein, but is rich in
     .alpha.G (18.5 kDa); (4) at pH 5, the most cationic fraction of
    human urinary protein constituted only about 4% of the total while
     the analogous fraction of rat urine, containing .alpha.G, contained 26% of
     total urinary protein; and (5) cationic (at pH 5.0) human
     urinary proteins included small amounts of proteins, e.g., .alpha.1-acid
     glycoprotein, and .alpha.1-microglobulin, which are products of the gene
     family coding for .alpha.G in rat. Thus, although humans excrete
     trace amounts of proteins similar to .alpha.G, the very low protein
     content of human urine, the relatively small proportion of
     cationic to total proteins, and the high MW of the most abundant
    human urinary proteins form a biological basis for suggesting that
    humans are not at risk for the type of fuel and solvent
    hydrocarbon-induced nephropathy, and the sequelae of such nephropathy,
    observed in male rats.
    Medical Descriptors:
     *hyaline droplet
     *kidney disease
     rat
    urine
    human cell
    animal cell
    human
    nonhuman
    male
    article
    priority journal
    Drug Descriptors:
     *alpha 1 microglobulin
     *orosomucoid
    protein
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(or